T-951 P.004

AMENDMENTS TO THE CLAIMS

Please amend claims 53 and 56 and cancel claim 59, as shown in the following listing of the claims:

- (currently amended) An ApoA-I agonist compound comprising: 53.
 - (i) a 18 to 22-residue peptide analogue that forms an amphipathic α-helix in the presence of lipids and that comprises formula (I):

 $Z_{1}-X_{1}-X_{2}-X_{3}-X_{4}-X_{5}-X_{6}-X_{7}-X_{8}-X_{9}-X_{10}-X_{11}-X_{12}-X_{13}-X_{14}-X_{15}-X_{16}-X_{17}-X_{18}-Z_{2}$ or a pharmaceutically acceptable salt thereof, wherein

X1 is Pro (P), Ala (A), Gly (G), Asn (N), Gln (Q) or D-pro (p);

X2 is an aliphatic residue;

 X_3 is Leu (L);

Feb-03-2006 01:04pm

From-

X4 is an acidic residue;

X₅ is Leu (L) or Phe (F);

X₆ is Leu (L) or Phe (F);

X₇ is a basic residue;

X₈ is an acidic residue;

X₉ is Leu (L) or Trp (W);

X₁₀ is Leu (L) or Trp (W);

X11 is an acidic residue or Asn (N);

X₁₂ is an acidic residue;

X₁₃ is Leu (L), Trp (W) or Phe (F);

X14 is a basic residue or Leu (L);

X₁₅ is Gln (Q) or Asn (N);X₁₆ is a basic residue;

 X_{17} is Leu (L);

X₁₈ is a basic residue;

 Z_1 is H_2N_- , or $RC(O)NR_-$;

Z₂ is -C(O)NRR, -C(O)OR or -C(O)OH or a salt theroof;

each R is independently -H, (C_1-C_6) alkyl, $(G_1 C_2-C_6)$ alkenyl, $(G_1 C_2-C_6)$ alkynyl, (C5-C20) aryl, (C6-C26) alkaryl, 5-20 membered heteroaryl or 6-26 membered alkheteroaryl or a 1 to 4-residue peptide or peptide analogue;

each " - " between residues X_1 through X_{18} independently designates an amide linkage, a substituted amide linkage, an isostere of an amide or an amide mimetic, wherein at least one "-" is a substituted amide linkage, an isostere of an amide or an amide mimetic;

Fab-03-2006 01:04pm

From-

P.005

(ii) a 15 to 21-residue peptide analogue according to formula (I) in which at least one and up to eight of residues X_1 , X_2 , X_3 , X_4 , X_5 , X_6 , X_7 , X_8 , X_9 , X_{10} , X_{11} , X_{12} , X_{13} , X_{14} , X_{15} , X_{16} , X_{17} and X_{18} are optionally deleted and wherein at least one "-" is a substituted amide linkage, an isostere of an amide or an amide mimetic; or

7346222928

- (iii) an 18 to 22-residue altered peptide analogue according to formula (I) in which at least one of residues X_1 , X_2 , X_3 , X_4 , X_5 , X_6 , X_7 , X_8 , X_9 , X_{10} , X_{11} , X_{12} , X_{13} , X_{14} , X_{15} , X_{16} , X_{17} and X_{18} is conservatively substituted and wherein at least one "-" is a substituted amide linkage, an isostere of an amide or an amide mimetic; or an N-terminally blocked form, a C-terminally blocked form or an N- and Cterminally blocked form of formula (I).
- (previously presented) The ApoA-I agonist compound of Claim 53 which exhibits at 54. least about 38% LCAT-activation activity as compared with human ApoA-I.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein at least 55. one "-" is a substituted amide linkage.
- (currently amended) The ApoA-I agonist compound of Claim 55 wherein the **56**. substituted amide linkage has the formula -C(O)NR-, where R is (C1-C6) alkyl, substituted (C_1 - C_6) alkyl, (C_1 C_2 - C_6) alkenyl, substituted (C_4 C_2 - C_6) alkenyl, (C_4 C_4 - C_6) alkenyl, (C_4 C_4 - C_6) alkenyl, (C_4 C_6 - C_6) alkenyl, (C_6 C_6 - C_6) alkenyl, (C_6 - C_6 - C_6) alkenyl, (C_6 - C_6 - C_6) alkenyl, (C_6 - C_6 - C_6 - C_6 - C_6 - $C_$ C₆) alkynyl, substituted (C_1 C_2 -C₆) alkynyl, (C_5 -C₂₀) aryl, substituted (C_5 -C₂₀) aryl, (C₆-C₂₆) alkaryl, substituted (C₆-C₂₆) alkaryl, 5-20 membered heteroaryl, substituted 5-20 membered heteroaryl, 6-26 membered alkheteroaryl, or substituted 6-26 membered alkheteroaryl.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the least 57. one "-" is an isostere of an amide.
- (previously presented) The ApoA-I agonist compound of Claim 57 wherein the 58. isostere of an amide is -CH2NH-, -CH2S-, CH2CH2-, -CH=CH- (cis and trans), -C(O)CH₂-, -CH(OH)CH₂-, or -CH₂SO-.
- (canceled). 59.

Fab-03-2006 01:04pm

T-951

P.006

- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the 60. peptide analogue exhibits 40% to 98% helicity in the presence of lipids.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the 61. peptide analogue comprises 40% to 70% hydrophobic residues.
- (previously presented) The ApoA-I agonist compound of Claim 61 wherein the 62. peptide analogue comprises 50% to 60% hydrophobic residues.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the mean 63. hydrophobic moment, $\langle \mu_H \rangle$, of the peptide analogue is 0.55 to 0.65.
- (previously presented) The ApoA-I agonist compound of Claim 63 wherein the mean 64. hydrophobic moment, $<\mu_H>$, of the peptide analogue is 0.58 to 0.62.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the 65. mean hydrophobicity, <H_o>, of the peptide analogue is -0.150 to -0.070.
- (previously presented) The ApoA-I agonist compound of Claim 65 wherein the mean 66. hydrophobicity, $\langle H_o \rangle$, of the peptide analogue is -0.130 to -0.050.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the mean 67. hydrophobicity of the hydrophobic face, $\langle H_o^{pho} \rangle$, of the peptide analogue is 0.90 to 1.20.
- (previously presented) The ApoA-I agonist compound of Claim 67 wherein the mean 68. hydrophobicity of the hydrophobic face, <H_o^{pho}>, of the peptide analogue is 0.95 to 1.10.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the pho 69. angle of the peptide analogue is 120° to 160°.
- (previously presented) The ApoA-I agonist compound of Claim 69 wherein the pho 70. angle of the peptide analogue is 130° to 150°.

Fab-03-2006 01:05pm

From-

T-951 P.007

- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the 71. peptide analogue has 3 to 5 positively charged amino acids.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the 72. peptide analogue has 3 to 5 negatively charged amino acids.
- (previously presented) The ApoA-I agonist compound of Claim 54 wherein the 73. peptide analogue has a net charge of -1, 0, or +1.
- (previously presented) An ApoA-I agonist-lipid complex comprising an ApoA-I 74. agonist compound and a lipid, wherein the ApoA-I agonist compound is a peptide analogue according to any one of claims 53-73.
- (previously presented) A pharmaceutical composition comprising an ApoA-I agonist 75. compound according to any one of claims 53-73 or an ApoA-I agonist-lipid complex according to claim 74, and a pharmaceutically acceptable carrier, excipient or diluent.
- (previously presented) A method of treating a subject suffering from a disorder 76. associated with dyslipidemia, said method comprising the step of administering to the subject an effective amount of the ApoA-I agonist compound of claim 53.
- (previously presented) The method of Claim 76 in which the disorder associated with *77*. dyslipidemia is hypercholesterolemia.
- (previously presented) The method of Claim 76 in which the disorder associated with 78. dyslipidemia is cardiovascular disease.
- (previously presented) The method of Claim 76 in which the disorder associated with 79. dyslipidemia is atherosclerosis.
- (previously presented) The method of Claim 76 in which the disorder associated with 80. dyslipidemia is restenosis.
- (previously presented) The method of Claim 76 in which the disorder associated with 81. dyslipidemia is HDL or ApoA-I deficiency.

- (previously presented) The method of Claim 76 in which the disorder associated with 82. dyslipidemia is hypertriglyceridemia.
- (previously presented) The method of Claim 76 in which the disorder associated with 83. dyslipidemia is metabolic syndrome.